# 002317112

# **Active Projects**

Following is a list of the principal investigators, or institutions, of projects under way or activated in the period since the previous Report, together with the respective project titles. Completed projects are listed in a later section.

# PRINCIPAL INVESTIGATOR OR INSTITUTION

# JOHN J. ALBERS, Ph.D., Research Associate Professor of Medicine, University of Washington School of Medicine, Seattle.

### HARRY N. ANTONIADES, Ph.D., Professor of Biochemistry, Harvard University School of Public Health, Boston.

- THOMAS M. AUNE, Ph.D., Adjunct in Immunology, The Jewish Hospital of St. Louis.
- BERNARD M. BABIOR, M.D., Ph.D., Professor of Medicine, New England Medical Center Hospital, Boston.
- LESLIE BAER, M.D., Associate Professor of Medicine, Columbia University College of Physicians & Surgeons, New York.
- BEA J. van den BERG, M.D., Research Pediatrician, Adjunct Professor in Biostatistics, University of California School of Public Health, Oakland.
- RICHARD J. BING, M.D., Professor of Medicine (emeritus), University of Southern California School of Medicine, Los Angeles; Visiting Associate in Biomedical Engineering, California Institute of Technology; Director of Cardiology and Intramural Medicine, Huntington Memorial Hospital, Pasadena, CA.
- DEBAJIT K. BISWAS, Ph.D., D.Sc.,
  Associate Professor of Oral Biology,
  Laboratory of Pharmacology, Harvard
  School of Dental Medicine, Boston.
- IRA B. BLACK, M.D., Professor and Chief, Division of Developmental Neurology, Cornell University Medical College, New York.
- PHYLLIS B. BLAIR, Ph.D., Professor of Immunology, University of California, Berkeley.

- High density lipoprotein quantitation
- Human platelet-derived growth factor (PDGF): relationship to human atherosclerosis
- Interferon—activation of suppressor T cell pathways
- Studies on the mechanism of activation of the respiratory burst in neutrophils
- Cigarette smoking in normotensive and hypertensive subjects: blood pressure, renin, aldosterone and catecholamine responses
- Pulmonary function tests in adolescents and their parents—an epidemiologic approach
- Cholesterol inhibition and carbon monoxide and atherosclerosis
- Lipoproteins and the arterial wall
- Effects of nicotine and benzo(a)pyrene on hormone production
- Nicotine and neuronal development
- Regulation of natural killer cell activity

# J. MARK BRAUGHLER, Ph.D., Assistant Professor of Pharmacology, Northeastern Ohio Universities College of Medicine, Rootstown.

EDWARD BRESNICK, Ph.D., Professor and Chairman, Department of Biochemistry, The University of Vermont College of Medicine, Burlington.

REBECCA BRYSON, Ph.D., Associate Professor of Psychology, San Diego State University, San Diego, CA.

VINCENZO BUONASSISI, M.D., Associate Research Biologist, The University of California at San Diego, La Jolla.

DAVID L. BUSBEE, Ph.D., Director, Genetics Center; Associate Professor of Biological Sciences, North Texas State University, Denton.

EDWARD J. CAMPBELL, M.D., Assistant Professor of Medicine, Washington University School of Medicine, St. Louis.

WILLIAM A. CARTER, M.D., Professor of Hematology and Medical Oncology, Hahnemann Medical College, Philadelphia.

ALBERT CASTRO, Ph.D., Director, Hormone Research Laboratory; Professor of Pathology and Medicine, University of Miami School of Medicine, Miami, FL.

FRANCIS C. CHAO, M.D., Ph.D., Senior Investigator, Center for Blood Research, Boston.

JAN F. CHLEBOWSKI, Ph.D., Assistant Professor of Biochemistry, Medical College of Virginia, Richmond.

CURT I. CIVIN, M.D., Assistant Professor of Oncology & Pediatrics, The Johns Hopkins Oncology Center, Baltimore.

CHARLES G. COCHRANE, M.D., Member, Department of Immunopathology, Scripps Clinic and Research Foundation, La Jolla, CA.

## PROJECT TITLE

The alteration of guanylate cyclase by nitric oxide

Expression of cytochrome P450c

Interactive effects of nicotine, testosterone and estradiol on weight change, food consumption and activity of male and female rats under high and low protein diets

Heparan sulfate proteoglycans and blood homeostatic mechanisms

The biochemical and physiological characteristics of a protein which specifically binds polycyclic aromatic hydrocarbons

Modulators of inflammatory cell proteolytic activity

The interplay of immunosurveillance and interferon induction in tumorigenesis

Nicotine in blood: detection by radioimmunoassay

Platelet activation and blood hypercoagulability

Calofimetric investigation of proteinase-\alpha\_2
macroglobulin interaction

Biochemistry and function of human granulopoietic antigens

Mediation systems in inflammatory lung disease

- BERNICE H. COHEN, Ph.D., Professor and Director, Human Genetics/Genetic Epidemiology Program, The Johns Hopkins University School of Hygiene and Public Health, Baltimore.
- ROBERT W. COLMAN, M.D., Professor of Medicine, Temple University School of Medicine, Philadelphia.
- GIDON CZAPSKI, M.Sc., Ph.D., Professor of Physical Chemistry, The Hebrew University, Jerusalem, Israel.
- ROBERT ECHT, Ph.D., Professor of Anatomy, Michigan State University, East Lansing.
- CARLTON K. ERICKSON, Ph.D., Professor of Pharmacology, The University of Texas College of Pharmacy, Austin.
- V. GENE ERWIN, Ph.D., Professor of Pharmacology; Dean; University of Colorado School of Pharmacy, Boulder.
- ALVAN R. FEINSTEIN, M.D., Professor of Medicine & Epidemiology, Yale University School of Medicine, New Haven, CT.
- JOSEPH D. FELDMAN, M.D., Immunopathologist, Scripps Clinic and Research Foundation, La Jolla, CA.
- THOMAS H. FINLAY, Ph.D., Associate Professor of Obstetrics and Gynecology, New York University Medical Center, New York.
- BIRGITTA FLODERUS-MYRRHED, Ph.D., Assistant Professor of Environmental Hygiene, The Karolinska Institute, Stockholm.
- JUDITH ANN FOSTER, Ph.D., Professor and Chairperson, Department of Biology, Sycracuse University, Syracuse, NY.
- JACK W. FRANKEL, Ph.D., Consultant in Medical Research, Veterans Administration Medical Center, Bay Pines, FL.
- ALLAN P. FREEDMAN, M.D., Assistant Professor of Medicine, Hahnemann Medical College, Philadelphia.

- Genetic-epidemiologic characteristics of smokers and nonsmokers
- Initiation of plasma coagulation and kinin forming systems in man
- On the toxicity of oxygen and superoxide ion: is superoxide toxic?
- The effects of hypoxic hypoxia, carbon monoxide and treatments influencing hypoxia toxicity on endocrine-like cells in airways of young rabbits and rabbit fetuses.
- Blood-brain monitoring of sustained nicotine levels in rats
- Actions of nicotine on isolated perfused mouse brain
- Effects of nicotine on neuropeptide secrection by intact mouse brain, a pharmacogenetic study
- Smoking, detection bias and primary lung cancer
- Effects of aging and lipids on the immune system
- Structure, properties and regulation of mouse plasma protease inhibitors
- Epidemiologic research on the Swedish twin registries
- Involvement of elastin in lung disease
- Smoking and lung cancer: diagnostic test to identify persons at high risk
- The effect of cigarette smoking on the alveolar clearance rate of inert dust particles in the human lung

- AARON E. FREEMAN, Ph.D., Staff Scientist, California Biomedical Research Foundation, La Jolla, CA.
- KJELL FUXE, M.D., Professor of Histology, The Karolinska Institute, Stockholm.
- MORTON GALDSTON, M.D., Associate Professor of Medicine, New York University Medical Center, New York.
- MICHAEL C. GEOKAS, M.D., Ph.D., Professor and Vice-Chairman, Department of Medicine, University of California School of Medicine, Davis.
- TERESA GESSNER, Ph.D., Associate Cancer Research Scientist, Health Research, Inc., Roswell Park Division, Ruffalo.
- JACQUES E. GIELEN, Ph.D.. Associate Professor, Laboratory of Medical Chemistry, Toxicology and Hygiene, Institute of Pathology, University of Liège, Liège, Belgium.
- GABRIEL C. GODMAN, M.D., Professor of Pathology, Columbia University College of Physicians & Surgeons, New York.
- WARREN M. GOLD, M.D., Professor of Medicine, Cardiovascular Research Institute, University of California, San Francisco.
- SIDNEY GOLDFISCHER, M.D., Professor of Pathology, Albert Einstein College of Medicine, The Bronx, NY.
- MAURICE GREEN, M.D., Director, Institute for Molecular Virology, St. Louis University Medical Center, St. Louis.
- MARK I. GREENE, Ph.D., Associate Professor of Pathology, Harvard Medical School, Boston.
- HIRA L. GURTOO, D.V.M., M.V. Sc., Ph.D., Associate Cancer Research Scientist, Department of Experimental Therapeutics, Roswell Park Memorial Institute, Buffalo.

- Rodent and human lung epithelial cell culture as a tool for carcinogenesis research in vitro
- Epithelial cell carcinogenesis in vitro
- Nicotine, catecholamines and neuroendocrine functions
- Biochemical basis of predisposition to chronic obstructive pulmonary disease
- Circulating pancreatic elastase 2 and emphysema in man
- Pharmacogenetics of conjugations and lung cancer risk
- Modulation of aryl hydrocarbon hydroxylase and epoxide hydratase in animal tissues and in cell culture by cigarette smoke condensate and other chemicals
- Biochemical mechanism(s) and qualitative and quantitative consequences of benzo(α) pyrene metabolism
- Cytoskeletal organization of the endothelial cell in regulation of shape contractility and sunface movement
- Effect of ozone on airway mast cells
- Extracellular matrix-cytochemistry and ultrastructure
- Amplification of human adenovirus transformation proteins in prokaryotic and eukaryotic cells
- Suppressor cells in syngeneic tumor immunity
- Role of genetics and polyaromatic hydrocarbon metabolism in human lung cancer

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# PRINCIPAL INVESTIGATOR OR INSTITUTION

- NOBUYOSHI HAGINO, M.D., Ph.D., Professor of Anatomy, University of Texas Health Science Center, San Antonio.
- CAROLINE B. HALL, M.D., Associate Professor of Pediatrics and Medicine, University of Rochester School of Medicine, Rochester, NY.
- LINDA M. HALL, Ph.D., Associate Professor of Genetics and Neuroscience, Albert Einstein College of Medicine of Yeshiva University, The Bronx, NY.
- PAUL HAMOSH, M.D., Associate Professor of Physiology and Biophysics, and Medicine, Georgetown University Schools of Medicine and Dentistry, Washington, D.C.
- NORMAN W. HEIMSTRA, Ph.D., Professor of Psychology; Director, Human Factors Laboratory, University of South Dakota, Vermillion.
- HERBERT B. HERSCOWITZ, Ph.D., Associate Professor of Microbiology, Georgetown University Schools of Medicine and Dentistry, Washington, D.C.
- ROBERT M. HOFFMAN, Ph.D., Assistant Professor of Pediatrics in Residence, University of California School of Medicine, La Jolla.
- JEROME L. HOJNACKI, Ph.D., Assistant Professor of Biological Sciences, University of Lowell, Lowell, MA.
- WAYNE HOSS, Ph.D., Associate Professor, Center for Brain Research, University of Rochester Medical Center, Rochester, NY.
- AARON JANOFF, Ph.D., Professor of Pathology, Health Sciences Center, State University of New York at Stony Brook, Stony Brook.
- MORRIS J. KARNOVSKY, M.B., B. Ch., Shattuck Professor of Pathological Anatomy, Harvard Medical School, Boston.
- INGEGARD M. KEITH, Ph.D., Assistant Professor of Anatomy, University of Wisconsin School of Veterinary Medicine, Madison.

- Nicotinic receptors of LHRH axon terminals in the median eminence
- Interrelationship of infectious lower respiratory tract disease in infancy, and host and environmental factors to later development of chronic lung disease
- Genetic differences in nicotine sensitivity in Drosophila melanogaster strains
- Cigarette smoke and lipoprotein remodeling by the lung
- Some behavioral aspects of smoking and smoking deprivation
- Effects of cigarette smoke exposure on developmental, cellular and molecular aspects of the immune response
- Methionine dependence, methylation and organic transformation
- Regulation of cellular oncogenes
- Nicotine-induced changes in primate high density lipoproteins
- Studies of nicotine interaction with blood cells
- Further studies on suppression of protease inhibition by cigarette smoke
- Immunologic assay of lung elastin degradation
- The molecular basis of pulmonary surfactant secretion by type II pneumocytes: studies in intact cells and a cell free system.
- Part I: Lung neuroendocrine cell innerva-
- Part II: Transplacental effect of smoking on lung neuroendocrine cells in the neonate

ROBERT W. KREILICK, Ph.D., Professor of Chemistry, University of Rochester, Rochester, NY.

KLAUS E. KUETTNER, Ph.D., Professor and Chairman; Department of Biochemistry, Rush College of Health Sciences and Rush Medical College, Rush-Presbyterian-St. Luke's Medical Center, Chicago.

LAWRENCE L. KUPPER, Ph.D., Associate Professor of Biostatistics, University of North Carolina School of Public Health, Chapel Hill.

ABEL LAJTHA, Ph.D., Director, New York State Research Institute for Neurochemistry and Drug Addiction, New York.

DON LAPENAS, M.D., Assistant Professor of Pathology, University of Vermont College of Medicine, Burlington.

E. CLINTON LAWRENCE, M.D., Assistant Professor of Medicine, Baylor College of Medicine, Houston.

PHILIP M. LE QUESNE, Ph.D., D.Sc., Professor of Chemistry, Northeastern University, Boston.

FABIAN J. LIONETTI, Ph.D., Research Professor of Biochemistry, Boston University School of Medicine, Boston

GESINA L. LONGENECKER, Ph.D.,

Associate Professor of Pharmacology,

University of South Alabama College

of Medicine, Mobile.

RONALD B. LUFTIG, Ph.D., Professor of Microbiology and Immunology, University of South Carolina School of Medicine, Columbia.

HENRY T. LYNCH, M.D., Professor and Chairman, Department of Preventive Medicine and Public Health, Creighton University School of Medicine, Omaha.

BRUCE A. MACHER, Ph.D., Assistant Professor of Pharmaceutical Chemistry, University of California, San Francisco.

ALAN C. McLAUGHLIN, Ph.D., Biophysicist, Brookhaven National Laboratory, Upton, NY.

### PROJECT TITLE

Investigations of the interaction of nicotine with membranes

Local regulation of normal and pathologic invasion

Regulation of proliferation of invasive cells

Verification of a statistical age-periodcohort analysis of lung cancer

Genetic basis for nicotine response

The association of inorganic dust deposition with pulmonary neoplasia in tobacco users

Effects of cigarette smoking on immunoglobulin production by human bronchial lymphocytes

Assay of oxygenated sterols in human blood vessels—a feasibility study

Phagocyte mediated injury to tissues

Studies of platelet and endothelial prostanoid production as possible cardiovascular risk indicators in smokers

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Interactions between RNA tumor viruses and chemical carcinogens

Genetic and biomarker studies of smoking-associated cancers

Chemistry and biology of complex carbohydrates

Interaction of divalent cations with model and biological membranes

- J. WISTER MEIGS, M.D., Clinical Professor of Epidemiology; Director, Connecticut Cancer Epidemiology Unit, Yale University School of Medicine, New Haven, CT.
- FERID MURAD, M.D., Ph.D., Professor of Medicine and Pharmacology, Stanford University, and Chief of Medicine, Palo Alto V.A. Hospital, Stanford, CA.
- JAY A. NADEL, M.D., Professor of Medicine, Physiology and Radiology, Cardiovascular Research Institute, University of California, San Francisco.
- DONALD J. NEUSON, Ph.D., Associate Professor of Chemistry, Clark University, Worcester, MA.
- HAROLD H. NEWBALL, M.D., Associate Professor of Medicine, The Johns Hopkins University School of Medicine, Baltimore.
- FRANZ OESCH, Ph.D., Professor of Pharmacology; Head, Section on Biochemical Pharmacology, University of Mainz, Mainz, West Germany.
- BENDICHT U. PAULI, D.V.M., Associate Professor of Pathology, Rush Presbyterian-St. Lukes Medical Center, Chicago.
- DENNIS R. PETERSEN, Ph.D., Assistant Professor of Pharmacology, University of Colorado School of Pharmacy, Boulder.
- JULIA M. POLAK, D.Sc., M.D., Senior Lecturer in Histopathology, Royal Postgraduate Medical School, Hammersmith Hospital, London.
- WILLIAM A. PRYOR, Ph.D., Boyd Professor of Chemistry, Louisiana State University, Baton Rouge.
- ILARI RANTASALO, M.D., Professor and Chairman, Department of Public Health Science, University of Helsinki, Helsinki, Finland.
- RONALD E. RASMUSSEN, Ph.D., Associate Adjunct Professor in Community and Environmental Medicine, University of California College of Medicine, Irvine.

### PROJECT TITLE

- Review of lung cancer in Connecticut, 1935-present
- Mechanism of nitric oxide activation of guanylate cyclase
- Role of cyclic GMP in smooth muscle relaxation
- Mechanisms of airway hyperinritability
- Spectroscopic studies of the interaction of cholinergic ligands with nicotinic receptor proteins
- The role of proteases and antiproteases in pulmonary emphysema
- Metabolic fate and toxicological significance of dihydrodiols derived from polycyclic aromatic hydrocarbons occurring in cigarette smoke
- Local regulation of tumor invasion by host-derived proteinase inhibitors
- Influences of genotype, sex and chronic cigarette smoking on nicotine and alcohol metabolism in mice
- Investigation of the role of regulatory peptides in human lung disease
- Free radical chemistry of cigarette smoke

- The Finnish Twin Cohort Follow-up Study
- The role of cell-specific toxins in mouse lung carcinogenesis

# EILEEN REMOLD-O'DONNELL, Ph.D., Principal Research Associate, Harvard Medical School; Investigator,

Center for Blood Research, Boston.

JOHN E. REPINE, M.D., Assistant Director, Webb-Waring Lung Institute; Associate Professor of Medicine, University of Colorado Health Sciences Center, Denver.

HERBERT Y. REYNOLDS, M.D., Professor of Medicine: Head, Pulmonary Section, Yale University School of Medicine, New Haven, CT.

VIRGINIA L. RICHMOND, Ph.D., Research Associate, Pacific Northwest Research Foundation, Seattle.

PETER M. ROSS, Ph.D., Research Associate, The Rockefeller University, New York.

UNA S. RYAN, Ph.D., Research Professor of Medicine, University of Miami School of Medicine, Miami, FL.

B. V. RAMA SASTRY, D.Sc., Ph.D., Professor of Pharmacology, Vander-bilt University School of Medicine, Nashville, TN.

ISHAIAHU SCHECHTER, PHD., Senior Lecturer in Biochemistry, The George S. Wise Faculty for Life Sciences, Tel Aviv University, Tel Aviv, Israel.

GERALD SHKLAR, D.D.S., Charles A. Brackett Professor of Oral Pathology; Head, Department of Oral Medicine and Oral Pathology, Harvard School of Dental Medicine, Boston.

ROBERT JI SKLAREW, Ph.D., Research Associate Professor of Pathology, New Manual University Research Service; Goldwater Memorial Hospital, Roosevelt Island, New York.

HANOCH SLOR, Ph.D., Associate Professor of Human Genetics, Sackler School of Medicine, Tel Aviv University, Tel Aviv, Israel.

DENNIS M. SMITH, Ph.D., Assistant Professor of Biological Sciences, Wel-lesley College, Wellesley, MA.

### PROJECT TITLE

Purification and functional analysis of elastase from guinea pig macrophages

Basic mechanisms of lung injury from inhaled oxidants

Respiratory secretions in pulmonary carcinoma: secretory component of immunoglobulin-A as an early marker of epithelial dysfunction

Markers of epithelial cell dysfunction in respiratory secretions of smokers

Elastic fiber microfibrillar protein struc-

DNA damage and selective maintenance of animal genes

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Interactions of hormones with cells of the pulmonary vascular wall

Influence of nicotine on the release of acetylcholine in the human placenta and its implications on the fetal growth

Effect of thiols and disulfides on cholesterol metabolism

Oral carcinogenesis, vitamin A and ret-

Cytokinetics of heteroploid subpopulations by imaging

The use of specific antibodies to monitor the formation and removal of benzo-(α)pyrene adducts from DNA of damaged human cells

Autonomic control of pulmonary surfactant in the adult lung

- TIMOTHY A. SPRINGER, Ph.D., Assistant Professor of Pathology, Harvard Medical School, Boston.
- ERIC J. STANBRIDGE, Ph.D., Associate Professor of Microbiology, University of California, Irvine.
- DANIEL STEINBERG, M.D., Ph.D., Professor of Medicine; Head, Division of Metabolic Disease, The University of California at San Diego, La Jolla.
- THOMAS P. STOSSEL, M.D., Chief, Medical Oncology Unit, Massachusetts General Hospital, Boston.
- D. LANSING TAYLOR, Ph.D., Associate Professor of Biology, Harvard University, Cambridge, MA.
- JAMES TRAVIS, Ph.D., Professor of Biochemistry, The University of Georgia, Athens.
- EMIL R. UNANUE, M.D., Mallinckrodt Professor of Immunopathology, Harvard Medical School, Boston.
- STEPHEN F. VATNER, M.D., Associate Professor of Medicine, Harvard Medical School, New England Regional primate Research Center, Southboro. MA.; Associate in Medicine, Peter Bent Brigham Hospital, Boston.
- PETER N. WALSH. Ph.D., Professor of Medicine, Temple University School of Medicine, Philadelphia.
- GEORGE WEINBAUM, PH.D., Bioscientist, Pulmonary Disease Section, Albert Einstein Medical Center, Philadelphia.
- I. BERNARD WEINSTEIN, M.D., Professor of Medicine and Environmental Sciences, Columbia University, New York.
- SIGMUND A. WEITZMAN, M.D., Assistant Professor of Medicine, Hematology—Oncology Unit, Massachusetts General Hospital, Boston.
- AKE WENNMALM, M.D., Associate Professor of Clinical Physiology at Karolinska Institute, Huddinge Hospital, Huddinge, Sweden.
- JOHN T. WILSON, Ph.D., Assistant Professor of Cell and Molecular Biology, Medical College of Georgia, Augusta.

- Studies of macrophage subpopulations and differentiation using monoclonal antibodies
- Transfer of specific individual human chromosomes to recipient cells
- Arterial degradation of low density lipoproteins in vivo
- Functional anatomy of the lung macrophage
- Chemotaxis of macrophages
- Proteolytic enzymes and inhibitors in emphysema
- Physiopathology of normal and activated macrophages
- Direct effects of nicotine on brain circulation
- Interaction of platelets with coagulation factors IX and X
- Bronchioalveolar lavage of human smokers and nonsmokers: studies on cellchemotaxis, enzyme release and cellular ultrastructure
- Development of monoclonal antibodies to carcinogen-DNA adducts
- Studies of phagocyte—induced mutation
- Nicotine as inhibitor of prostaglandin formation: localization of the inhibitory step and characterization of the cardiovascular implications
- The isolation and expression of human α-1-antitrypsin gene sequences through molecular cloning

BRUCE A. WODA, M.D., Associate Professor of Pathology, University of Massachusetts, Worcester.

STANLEY YACHNIN, M.D., Professor of Medicine and Chief, Section of Hematology/Oncology, The University of Chicago Medical Center, Chicago.

# PROJECT TITLE

Effects of smoking on the inherent interferon levels in control and cancer patients: a pilot study

Cell surface membranes, integral membrane proteins and cytoskeleton in lymphocytes from: (1) young and old rats; (2) spontaneous rat lymphomas

Models for the pathogenesis of atherosclerosis: A) biological effects of oxygenated sterol compounds; B) mevalonic acid and cholesterol biosynthesis and the biosynthesis and regulation of cell growth